Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

1. (Currently Amended) A method of <u>increasing rate of skeletal repair in a mammal having a bone implant or bone transplant by stimulating osteoblast-mediated growth of new bone at the site of the transplant or <u>implant</u>, in a mammal comprising administering to a mammal in need thereof the mammal at the site of the implant or <u>transplant</u>, in an immobilized, slow release form, a therapeutically effective amount of a compound having the formula:</u>

$$Y_2O^{W}$$
 R_{6}
 R_{7}
 R_{7}

where Y_1 and Y_2 , which may be the same or different, are each selected from the group consisting of hydrogen and a hydroxy-protecting group, where R_{11} and R_{12} are each hydrogen or taken together are a methylene group, where R_6 and R_7 , which may be the same or different, are each selected from the group consisting of hydrogen, alkyl, hydroxyalkyl, fluoroalkyl, hydroxy and alkoxy, with the proviso that R_6 and R_7 cannot both be hydrogen, or R_6 and R_7 when taken together may represent the group -(CH₂)_x-

where X is an integer from 2 to 5, or R_6 and R_7 when taken together may represent the group = CR_8R_9 where R_8 and R_9 , which may be the same or different, are each selected from the group consisting of hydrogen, alkyl, hydroxyalkyl, fluoroalkyl, hydroxy and alkoxy, or when taken together R_8 and R_9 may represent the group - $(CH_2)_x$ - where X is an integer from 2 to 5, and where the group R represents

where the stereochemical center (corresponding to C-20 in steroid numbering) may have the \underline{R} or \underline{S} configuration, (i.e. either the natural configuration about carbon 20 or the 20-epi configuration), and where Z is selected from Y, -OY, -CH₂OY,

-C≡CY and -CH=CHY, where the double bond may have the cis or trans geometry, and where Y is selected from hydrogen, methyl, -COR⁵ and a radical of the structure:

$$-(CH_2)_m$$
 $\stackrel{R^1}{-}C$ $\stackrel{R^2}{-}(CH_2)_n$ $\stackrel{R^3}{-}C$ $\stackrel{R^5}{-}$ $\stackrel{R^4}{-}$

where m and n, independently, represent the integers from 0 to 5, where R^1 is selected from hydrogen, deuterium, hydroxy, protected hydroxy, fluoro, trifluoromethyl, and C_{1-5} -alkyl, which may be straight chain or branched and, optionally, bear a hydroxy or protected-hydroxy substituent, and where each of R^2 , R^3 , and R^4 , independently, is selected from deuterium, deuteroalkyl, hydrogen, fluoro, trifluoromethyl and C_{1-5} alkyl, which may be straight-chain or branched, and optionally, bear a hydroxy or protected-hydroxy substituent, and where R^1 and R^2 , taken together, represent an oxo group, or an alkylidene group, $=CR^2R^3$, or the group $-(CH_2)_{p^-}$, where p is an integer from 2 to 5, and

where R^3 and R^4 , taken together, represent an oxo group, or the group - $(CH_2)_q$ -, where q is an integer from 2 to 5, and where R^5 represents hydrogen, hydroxy, protected hydroxy, or C_{1-5} alkyl and wherein any of the CH-groups at positions 20, 22, or 23 in the side chain may be replaced by a nitrogen atom, or where any of the groups - $CH(CH_3)$ -, - $(CH_2)_m$ -, - CR_1R_2 - or - $(CH_2)_n$ - at positions 20, 22, and 23, respectively, may be replaced by an oxygen or sulfur atom.

2.-7. (Cancelled)

- 8. (Original) The method of claim 1 wherein the compound is administered in a dosage of from $0.01\mu g$ to $50\mu g$ per day.
 - 9. (Original) The method of claim 1 wherein the mammal is a human.
- 10. (Original) The method of claim 1 wherein the compound administered is 2-methylene-19-nor-20(S)- 1α ,25-dihydroxyvitamin D₃ having the formula:

11. (Original) The method of claim 1 wherein the compound administered is an acylated derivative having the formula:

$$Y_2O^{m}$$
 R_{11}
 R_{12}
 OY_1

where Y^1 and Y^2 independently represent hydrogen or an acyl group, and with the proviso that R^5 is $-OY_3$ and Y_3 is selected from the group consisting of acyl or a hydrocarbyloxycarbonyl.

- 12. (Original) The method of claim 11 wherein the compound is a triacetate such that Y_1 , Y_2 and Y_3 and each CH_3CO -.
- 13. (Original) The method of claim 11 wherein the compound as a trihexanoate such that Y_1 , Y_2 and Y_3 are each $CH_3(CH_2)_4CO$ -.
- 14. (Original) The method of claim 11 wherein the compound is a trinonanoate such that Y_1 , Y_2 and Y_3 are each $CH_3(CH_2)_7CO$ -.
- 15. (Original) The method of claim 11 wherein the compound is a 25-acetate such that Y_1 and Y_2 are both hydrogen and Y_3 is CH_3CO -.
- 16. (Original) The method of claim 11 wherein the compound is 2-methylene-19-nor- 1α ,25(OH)₂-D₃-1,3,25-triacetate.

- 17. (Original) The method of claim 11 wherein the compound is 2-methylene-19-nor- 1α ,25(OH)₂-D₃-1,3,25-trihexanoate.
- 18. (Original) The method of claim 11 wherein the compound is 2-methylene-19-nor- 1α ,25(OH)₂-D₃-1,3,25-trinonanoate.
- 19. (Original) The method of claim 11 wherein the compound is 2-methylene-19-nor- 1α ,25(OH)₂-D₃-25-acetate.
- 20. (Original) The method of claim 1 wherein the compound administered is selected from the group consisting of:

$$Y_2OW$$
 R_0
 R_0
 R_0
 R_0

where Y_1 , Y_2 , R_{11} , R_{12} and R are as defined in claim 1 and R_8 and R_9 , which may be the same or different, are each selected from the group consisting of hydrogen, alkyl, hydroxyalkyl and fluoroalkyl, or, when taken together represent the group -(CH_2)_X- where X is an integer from 2 to 5.

21. (Original) The method of claim 1 wherein the compound administered is selected from the group consisting of:

$$Y_{2}O^{W}$$

$$X_{10}$$

$$X_{10}$$

$$X_{10}$$

$$X_{10}$$

where Y_1 , Y_2 , R_{11} and R_{12} and R are as defined in claim 1 and R_{10} is selected from the group consisting of alkyl, hydroxyalkyl and fluoroalkyl.

22. (Original) The method of claim 1 wherein the compound administered is selected from the group consisting of:

$$\begin{array}{c|c} R \\ \hline \\ H \\ \hline \\ Y_2O_{11} \\ \hline \\ R_6 \\ \hline \\ R_7 \\ \end{array}$$

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where Y_1 , Y_2 , R_{11} , R_{12} , R_6 , R_7 and R are as defined in claim 1 with the proviso that R^5 is - OY_3 and Y_3 is selected from the group consisting of an acyl or a hydrocarbyloxycarbonyl.

23.-28. (Cancelled)

29. (New) A method of increasing rate of skeletal repair in a mammal having a bone implant or bone transplant by stimulating osteoblast-mediated growth of new bone at the site of the transplant or implant, comprising administering to the mammal at the site of the implant or transplant, in an immobilized form, a therapeutically effective amount of a compound having the formula:

$$Y_2O^{WW}$$
 R_6
 R_7
 R_{12}
 R_{12}
 R_{12}
 R_{12}

where Y_1 and Y_2 , which may be the same or different, are each selected from the group consisting of hydrogen and a hydroxy-protecting group, where R_{11} and R_{12} are each hydrogen or taken together are a methylene group, where R_6 and R_7 , which may be the same or different, are each selected from the group consisting of hydrogen, alkyl, hydroxyalkyl, fluoroalkyl, hydroxy and alkoxy, with the proviso that R_6 and R_7 cannot both be hydrogen, or R_6 and R_7 when taken together may represent the group -(CH_2)_x-where X is an integer from 2 to 5, or R_6 and R_7 when taken together may represent the group = CR_8R_9 where R_8 and R_9 , which may be the same or different, are each selected from the group consisting of hydrogen, alkyl, hydroxyalkyl, fluoroalkyl, hydroxy and

alkoxy, or when taken together R_8 and R_9 may represent the group -(CH_2)_x- where X is an integer from 2 to 5, and where the group R represents

where the stereochemical center (corresponding to C-20 in steroid numbering) may have the \underline{R} or \underline{S} configuration, (i.e. either the natural configuration about carbon 20 or the 20-epi configuration), and where Z is selected from Y, -OY, -CH₂OY,

-C≡CY and -CH=CHY, where the double bond may have the cis or trans geometry, and where Y is selected from hydrogen, methyl, -COR⁵ and a radical of the structure:

$$-(CH_2)_m$$
 $-C$ $-(CH_2)_n$ $-C$ $-R^5$ R^4

where m and n, independently, represent the integers from 0 to 5, where R^1 is selected from hydrogen, deuterium, hydroxy, protected hydroxy, fluoro, trifluoromethyl, and C_{1-5} -alkyl, which may be straight chain or branched and, optionally, bear a hydroxy or protected-hydroxy substituent, and where each of R^2 , R^3 , and R^4 , independently, is selected from deuterium, deuteroalkyl, hydrogen, fluoro, trifluoromethyl and C_{1-5} alkyl, which may be straight-chain or branched, and optionally, bear a hydroxy or protected-hydroxy substituent, and where R^1 and R^2 , taken together, represent an oxo group, or an alkylidene group, $=CR^2R^3$, or the group $-(CH_2)_p$ -, where p is an integer from 2 to 5, and where R^3 and R^4 , taken together, represent an oxo group, or the group $-(CH_2)_q$ -, where q is an integer from 2 to 5, and where R^5 represents hydrogen, hydroxy, protected hydroxy,

or C_{1-5} alkyl and wherein any of the CH-groups at positions 20, 22, or 23 in the side chain may be replaced by a nitrogen atom, or where any of the groups -CH(CH₃)-, - $(CH_2)_m$ -, -CR₁R₂- or - $(CH_2)_n$ - at positions 20, 22, and 23, respectively, may be replaced by an oxygen or sulfur atom.

30. (New) The method of claim 29 wherein the compound administered is 2-methylene-19-nor-20(S)- 1α ,25-dihydroxyvitamin D₃ having the formula:

31. (New) A method of increasing rate of skeletal repair in a mammal having a bone implant or bone transplant by stimulating osteoblast-mediated growth of new bone at the site of the transplant or implant, comprising administering to the mammal at the site of the implant or transplant, in slow release form, a therapeutically effective amount of a compound having the formula:

$$Y_2O^{W}$$

$$R_{6}$$

$$R_{7}$$

$$R_{11}$$

$$R_{12}$$

$$OY_1$$

where Y_1 and Y_2 , which may be the same or different, are each selected from the group consisting of hydrogen and a hydroxy-protecting group, where R_{11} and R_{12} are each hydrogen or taken together are a methylene group, where R_6 and R_7 , which may be the same or different, are each selected from the group consisting of hydrogen, alkyl, hydroxyalkyl, fluoroalkyl, hydroxy and alkoxy, with the proviso that R_6 and R_7 cannot both be hydrogen, or R_6 and R_7 when taken together may represent the group -(CH_2)_x-where X is an integer from 2 to 5, or R_6 and R_7 when taken together may represent the group = CR_8R_9 where R_8 and R_9 , which may be the same or different, are each selected from the group consisting of hydrogen, alkyl, hydroxyalkyl, fluoroalkyl, hydroxy and alkoxy, or when taken together R_8 and R_9 may represent the group -(CH_2)_x- where X is an integer from 2 to 5, and where the group R represents

where the stereochemical center (corresponding to C-20 in steroid numbering) may have the \underline{R} or \underline{S} configuration, (i.e. either the natural configuration about carbon 20 or the 20-epi configuration), and where Z is selected from Y, -OY, -CH₂OY,

-C≡CY and -CH=CHY, where the double bond may have the cis or trans geometry, and where Y is selected from hydrogen, methyl, -COR⁵ and a radical of the structure:

$$-(CH_2)_m$$
 R^1 R^2 $(CH_2)_n$ C^3 R^5

where m and n, independently, represent the integers from 0 to 5, where R^1 is selected from hydrogen, deuterium, hydroxy, protected hydroxy, fluoro, trifluoromethyl, and $C_{1.5}$ -alkyl, which may be straight chain or branched and, optionally, bear a hydroxy or protected-hydroxy substituent, and where each of R^2 , R^3 , and R^4 , independently, is selected from deuterium, deuteroalkyl, hydrogen, fluoro, trifluoromethyl and C_{1-5} alkyl, which may be straight-chain or branched, and optionally, bear a hydroxy or protected-hydroxy substituent, and where R^1 and R^2 , taken together, represent an oxo group, or an alkylidene group, = CR^2R^3 , or the group - $(CH_2)_p$ -, where p is an integer from 2 to 5, and where R^3 and R^4 , taken together, represent an oxo group, or the group - $(CH_2)_q$ -, where q is an integer from 2 to 5, and where R^5 represents hydrogen, hydroxy, protected hydroxy, or C_{1-5} alkyl and wherein any of the CH-groups at positions 20, 22, or 23 in the side chain may be replaced by a nitrogen atom, or where any of the groups - $CH(CH_3)$ -, - $(CH_2)_m$ -, - CR_1R_2 - or - $(CH_2)_n$ - at positions 20, 22, and 23, respectively, may be replaced by an oxygen or sulfur atom.

32. (New) The method of claim 31 wherein the compound administered is 2-methylene-19-nor-20(S)- 1α ,25-dihydroxyvitamin D₃ having the formula: